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N-Ac-Sar-Gly-Val-D-Ile-Thr-Gln-Ile-Arg-ProNHCH<sub>2</sub>CH<sub>3</sub>, or a pharmaceutically acceptable salt, ester, solvate, or prodrug thereof, and a pharmaceutically acceptable carrier.

32 (New). A composition for the treatment of a disease selected from cancer, arthritis, psoriasis, angiogenesis of the eye associated with infection or surgical intervention, macular degeneration, and diabetic retinopathy comprising N-Ac-Sar-Gly-Val-D-alloIle-Ser-Ser-Ile-Arg-ProNHCH<sub>2</sub>CH<sub>3</sub>, or a pharmaceutically acceptable salt, ester, solvate, or prodrug thereof and a pharmaceutically acceptable carrier.

#### REMARKS

Claims 1-17 were pending in the subject application. After entry of this amendment, claims 1-14, 16, and 18-32 will be pending.

The amendment accompanying these remarks cancels claims 15 and 17 as being outside the scope of elected Group 1 and amends claim 1 to recite the amino acyl residue "glycyl" as option (13) under A<sub>2</sub>, to remove variables in claim 1 which are outside the scope of Group 1, and to correct a minor typographical error.

Support for the recitation of glycyl in claim 1 is found on page 19, lines 15-19 and in Examples 1-152, 161-334, 337 (in part), 339 (in part), and 340-356 in the specification.

#### CONCLUSION

This amendment addresses the issues raised in the Official action and places the claims in condition for allowance. Entry of the proposed amendment and allowance of Claims 1-14, 16, and 18-32 is respectfully requested.



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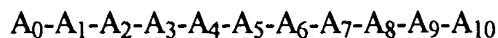
ABBOTT LABORATORIES  
Telephone: (847) 937-3810  
Facsimile: (847) 938-2623

Respectfully submitted,  
J. Henkin, et al.

*B. Gregory Donner*  
B. Gregory Donner  
Registration No. 34,580  
Attorney for Applicants

## VERSION OF CLAIMS WITH MARKINGS TO SHOW CHANGES MADE

1 (Amended). A compound of the formula:



or a pharmaceutically acceptable salt, ester, solvate or prodrug thereof, wherein:

$A_0$  is [hydrogen or] an acyl group selected from:

- (1)  $R-(CH_2)_n-C(O)-$ ; wherein  $n$  is an integer from 0 to 8 and  $R$  is selected from hydroxyl; methyl; N-acetylamino; methoxyl; carboxyl; cyclohexyl optionally containing a one or two double bonds and optionally substituted with one to three hydroxyl groups; and a 5- or 6-membered ring aromatic or nonaromatic ring optionally containing one or two heteroatoms selected from nitrogen, oxygen, and sulfur, wherein the ring is optionally substituted with a moiety selected from alkyl, alkoxy, and halogen; and
- (2)  $R^1-CH_2CH_2-(OCH_2CH_2O)_p-CH_2-C(O)-$ ; wherein  $R^1$  is selected from hydrogen, alkyl, and N-acetylamino, and  $p$  is an integer from 1 to 8;

$A_1$  is an amino acyl residue selected from:

- (1) alanyl,
- (2) asparaginyl,
- (3) citrullyl,
- (4) glutaminyl,
- (5) glutamyl,
- (6) N-ethylglycyl,
- (7) methionyl,
- (8) N-methylalanyl,
- (9) prolyl,
- (10) pyro-glutamyl,
- (11) sarcosyl,
- (12) seryl,

- (13) threonyl,
- (14)  $\text{-HN-(CH}_2\text{)}_q\text{-C(O)-}$ , wherein q is 1 to 8, and
- (15)  $\text{-HN-CH}_2\text{CH}_2\text{-(OCH}_2\text{CH}_2\text{O)}_r\text{-CH}_2\text{-C(O)-}$ , wherein r is 1 to 8;

A<sub>2</sub> is an amino acyl residue selected from:

- (1) alanyl,
- (2) asparaginylyl,
- (3) aspartyl,
- (4) glutaminylyl,
- (5) glutamyl,
- (6) leucyl,
- (7) methionyl,
- (8) phenylalanyl,
- (9) prolyl,
- (10) seryl,
- (11)  $\text{-HN-(CH}_2\text{)}_q\text{-C(O)-}$ , wherein q is 1 to 8, **[and]**
- (12)  $\text{-HN-CH}_2\text{CH}_2\text{-(OCH}_2\text{CH}_2\text{O)}_r\text{-CH}_2\text{-C(O)-}$ , wherein r is 1 to 8[;], **and**
- (13) glycyl;**

A<sub>3</sub> is an amino acyl residue selected from:

- (1) alanyl,
- (2) asparaginylyl,
- (3) citrullyl,
- (4) cyclohexylalanyl,
- (5) cyclohexylglycyl,
- (6) glutaminylyl,
- (7) glutamyl,
- (8) glycyl,
- (9) isoleucyl,
- (10) leucyl,
- (11) methionyl,
- (12) norvalyl,
- (13) phenylalanyl,
- (14) seryl,
- (15) *t*-butylglycyl,
- (16) threonyl,

- (17) valyl,
- (18) penicillaminy, and
- (19) cystyl;

A<sub>4</sub> is an amino acyl residue of L or D configuration selected from:

- (1) allo-isoleucyl,
- (2) glycyl,
- (3) isoleucyl,
- (4) prolyl,
- (5) dehydroleucyl,
- (6) D-alanyl,
- (7) D-3-(naphth-1-yl)alanyl,
- (8) D-3-(naphth-2-yl)alanyl,
- (9) D-(3-pyridyl)-alanyl,
- (10) D-2-aminobutyryl,
- (11) D-allo-isoleucyl,
- (12) D-allo-threonyl,
- (13) D-allylglycyl,
- (14) D-asparaginy, and
- (15) D-aspartyl,
- (16) D-benzothienyl,
- (17) D-3-(4,4'-biphenyl)alanyl,
- (18) D-chlorophenylalanyl,
- (19) D-3-(3-trifluoromethylphenyl)alanyl,
- (20) D-3-(3-cyanophenyl)alanyl,
- (21) D-3-(3,4-difluorophenyl)alanyl,
- (22) D-citrullyl,
- (23) D-cyclohexylalanyl,
- (24) D-cyclohexylglycyl,
- (25) D-cystyl,
- (26) D-cystyl(*S*-*t*-butyl),
- (27) D-glutaminy, and
- (28) D-glutamyl,
- (29) D-histidyl,
- (30) D-homoisoleucyl,

- (31) D-homophenylalanyl,
- (32) D-homoseryl,
- (33) D-isoleucyl,
- (34) D-leucyl,
- (35) D-lysyl(N-epsilon-nicotinyl),
- (36) D-lysyl,
- (37) D-methionyl,
- (38) D-neopentylglycyl,
- (39) D-norleucyl,
- (40) D-norvalyl,
- (41) D-ornithyl,
- (42) D-penicillaminyl,
- (43) D-penicillaminyl(acetamidomethyl),
- (44) D-penicillaminyl(*S*-benzyl),
- (45) D-phenylalanyl,
- (46) D-3-(4-aminophenyl)alanyl,
- (47) D-3-(4-methylphenyl)alanyl,
- (48) D-3-(4-nitrophenyl)alanyl,
- (49) D-3-(3,4-dimethoxyphenyl)alanyl,
- (50) D-3-(3,4,5-trifluorophenyl)alanyl,
- (51) D-prolyl,
- (52) D-seryl,
- (53) D-seryl(*O*-benzyl),
- (54) D-*t*-butylglycyl,
- (55) D-thienylalanyl,
- (56) D-threonyl,
- (57) D-threonyl(*O*-benzyl),
- (58) D-tryptyl,
- (59) D-tyrosyl(*O*-benzyl),
- (60) D-tyrosyl(*O*-ethyl),
- (61) D-tyrosyl, and
- (62) D-valyl;

A<sub>5</sub> is an amino acyl residue of L or D configuration selected from:

- (1) alanyl,

- (2) (3-pyridyl)alanyl,
- (3) 3-(naphth-1-yl)alanyl,
- (4) 3-(naphth-2-yl)alanyl,
- (5) allo-threonyl,
- (6) allylglycyl,
- (7) glutaminyl,
- (8) glycyl,
- (9) histidyl,
- (10) homoseryl,
- (11) isoleucyl,
- (12) lysyl(N-epsilon-acetyl),
- (13) methionyl,
- (14) norvalyl,
- (15) octylglycyl,
- (16) ornithyl,
- (17) 3-(4-hydroxymethylphenyl)alanyl,
- (18) prolyl,
- (19) seryl,
- (20) threonyl,
- (21) tryptyl,
- (22) tyrosyl,
- (23) D-allo-threonyl,
- (24) D-homoseryl,
- (25) D-seryl,
- (26) D-threonyl,
- (27) penicillaminyl, and
- (28) cystyl;

A<sub>6</sub> is an amino acyl residue of L or D configuration selected from:

- (1) alanyl,
- (2) 3-(naphth-1-yl)alanyl,
- (3) 3-(naphth-2-yl)alanyl,
- (4) (3-pyridyl)alanyl,
- (5) 2-aminobutyryl,
- (6) allylglycyl,

- (7) arginyl,
- (8) asparaginyl,
- (9) aspartyl,
- (10) citrullyl,
- (11) cyclohexylalanyl,
- (12) glutaminyl,
- (13) glutamyl,
- (14) glycyl,
- (15) histidyl,
- (16) homoalanyl,
- (17) homoleucyl,
- (18) homoseryl,
- (19) isoleucyl,
- (20) leucyl,
- (21) lysyl(N-epsilon-acetyl),
- (22) lysyl(N-epsilon-isopropyl),
- (23) methionyl(sulfone),
- (24) methionyl(sulfoxide),
- (25) methionyl,
- (26) norleucyl,
- (27) norvalyl,
- (28) octylglycyl,
- (29) phenylalanyl,
- (30) 3-(4-carboxyamidophenyl)alanyl,
- (31) propargylglycyl,
- (32) seryl,
- (33) threonyl,
- (34) tryptyl,
- (35) tyrosyl,
- (36) valyl,
- (37) D-3-(naphth-1-yl)alanyl,
- (38) D-3-(naphth-2-yl)alanyl,
- (39) D-glutaminyl,
- (40) D-homoseryl,

- (41) D-leucyl,
- (42) D-norvalyl,
- (43) D-seryl,
- (44) penicillaminyl, and
- (45) cystyl;

A<sub>7</sub> is an amino acyl residue of L or D configuration selected from:

- (1) alanyl,
- (2) allylglycyl,
- (3) aspartyl,
- (4) citrullyl,
- (5) cyclohexylglycyl,
- (6) glutamyl,
- (7) glycyl,
- (8) homoseryl,
- (9) isoleucyl,
- (10) allo-isoleucyl
- (11) leucyl,
- (12) lysyl(N-epsilon-acetyl),
- (13) methionyl,
- (14) 3-(naphth-1-yl)alanyl,
- (15) 3-(naphth-2-yl)alanyl,
- (16) norvalyl,
- (17) phenylalanyl,
- (18) prolyl,
- (19) seryl,
- (20) *t*-butylglycyl,
- (21) tryptyl,
- (22) tyrosyl,
- (23) valyl,
- (24) D-allo-isoleucyl,
- (25) D-isoleucyl,
- (26) penicillaminyl, and
- (27) cystyl;

A<sub>8</sub> is an amino acyl residue selected from:



- (1) 2-amino-4-[(2-amino)-pyrimidinyl]butanoyl,
- (2) alanyl(3-guanidino),
- (3) alanyl[3-pyrrolidinyl(2-N-amidino)],
- (4) alanyl[4-piperidinyl(N-amidino)],
- (5) arginyl,
- (6) arginyl(N<sup>G</sup>N<sup>G'</sup>diethyl),
- (7) citrullyl,
- (8) 3-(cyclohexyl)alanyl(4-N'-isopropyl),
- (9) glycyl[4-piperidinyl(N-amidino)],
- (10) histidyl,
- (11) homoarginyl,
- (12) lysyl,
- (13) lysyl(N-epsilon-isopropyl),
- (14) lysyl(N-epsilon-nicotinyl),
- (15) norarginyl,
- (16) ornithyl(N-delta-isopropyl),
- (17) ornithyl(N-delta-nicotinyl),
- (18) ornithyl[N-delta-(2-imidazoliny)],
- (19) [(4-amino(N-isopropyl)methyl)phenyl]alanyl,
- (20) 3-(4-guanidinophenyl)alanyl, and
- (21) 3-(4-amino-N-isopropylphenyl)alanyl;

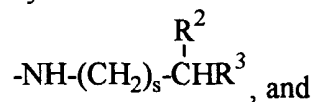
A<sub>9</sub> is an amino acyl residue of L or D configuration selected from:

- (1) 2-amino-butyryl,
- (2) 2-amino-isobutyryl,
- (3) homoprolyl,
- (4) hydroxyprolyl,
- (5) isoleucyl,
- (6) leucyl,
- (7) phenylalanyl,
- (8) prolyl,
- (9) seryl,
- (10) *t*-butylglycyl,
- (11) 1,2,3,4-tetrahydroisoquinoline-3-carbonyl,
- (12) threonyl,

- (13) valyl,
- (14) D-alanyl, and
- (15) D-prolyl; and

A<sub>10</sub> is a hydroxyl group or an amino acid amide is selected from:

azaglycylamide,  
 D-alanylamine,  
 D-alanylethylamine,  
 glycylamine,  
 glycylethylamine,  
 sarcosylamine,  
 serylamine,  
 D-serylamine,  
 a group represented by the formula



a group represented by the formula  $-\text{NH}-\text{R}^4$ ;

wherein:

s is an integer selected from 0 to 8,

R<sup>2</sup> is selected from hydrogen, alkyl, and a 5- to 6-membered cycloalkyl ring;

R<sup>3</sup> is selected from hydrogen, hydroxy, alkyl, phenyl, alkoxy, and a 5- to 6-membered ring optionally containing from one to two heteroatoms selected from oxygen, nitrogen, and sulfur, provided that s is not zero when R<sup>3</sup> is hydroxy or alkoxy; and

R<sup>4</sup> is selected from hydrogen, hydroxy, and a 5- to 6-membered cycloalkyl ring.